

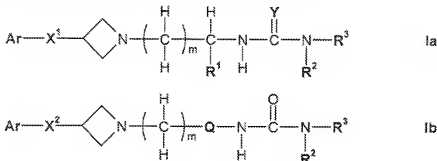
Amendments to the Claims

This listing of claims will replace all prior versions, and listings of claims in the specification:

Listing of Claims

Claims 1 -10. (Cancelled)

Claim 11. (Currently Amended) A compound of formula Ia or Ib



or its pharmaceutically acceptable salts in free or salt form, where

Ar is phenyl optionally substituted by one or more substituents selected from halogen,

C₁-C₈-alkyl, cyano or nitro;

X¹ is -S-, -S(=O)- or -S(=O)₂-;

X² is -C(=O)-, -O-, -CH₂-, -S-, -S(=O)- or -S(=O)₂-;

m is 1, 2, 3 or 4;

R¹ is hydrogen or C₁-C₈-alkyl optionally substituted by hydroxy, C₁-C₈-alkoxy, acyloxy, halogen, carboxy, C₁-C₈-alkoxycarbonyl, -N(R⁶)R⁵, -CON(R⁶)R⁵ or by a monovalent cyclic organic group having 3 to 15 atoms in the ring system;

Q has the formula



where R^a is C₁-C₈-alkylene,

or Q is -C(R^b)(R^c)- where R^b and R^c are independently C₁-C₈-alkyl

or R^b and R^c together form a C₃-C₁₀-cycloalkyl;

Y is oxygen or sulfur;

R² is hydrogen, C₁-C₈-alkyl or C₃-C₁₀-cycloalkyl and R³ is C₁-C₈-alkyl substituted by phenyl, phenoxy, acyloxy or naphthyl, or R³ is C₃-C₁₀-cycloalkyl optionally having a benzo group fused thereto, a heterocyclic group having 5 to 11 ring atoms of which 1 to 4 are hetero atoms, phenyl

or naphthyl, said phenyl, phenoxy or naphthyl groups being optionally substituted by one or more substituents selected from halogen, cyano, hydroxy, acyl, nitro, $-\text{SO}_2\text{NH}_2$, $\text{C}_1\text{-C}_6\text{-alkyl}$ optionally substituted by $\text{C}_1\text{-C}_6\text{-alkoxy}$, $\text{C}_1\text{-C}_6\text{-haloalkyl}$, $\text{C}_1\text{-C}_6\text{-alkoxy}$, $\text{C}_1\text{-C}_6\text{-haloalkoxy}$, $\text{C}_1\text{-C}_6\text{-alkylthio}$, $-\text{SO}_2\text{-C}_1\text{-C}_6\text{-alkyl}$, $\text{C}_1\text{-C}_6\text{-alkoxycarbonyl}$, $\text{C}_1\text{-C}_6\text{-acylamino}$ optionally substituted on the nitrogen atom by $\text{C}_1\text{-C}_6\text{-alkyl}$, $\text{C}_1\text{-C}_6\text{-alkylamino}$, aminocarbonyl, $\text{C}_1\text{-C}_6\text{-alkylamino-carbonyl}$, $\text{di}(\text{C}_1\text{-C}_6\text{-alkyl})\text{amino}$, $\text{di}(\text{C}_1\text{-C}_6\text{-alkyl})\text{aminocarbonyl}$, $\text{di}(\text{C}_1\text{-C}_6\text{-alkyl})\text{aminocarbonyl-methoxy}$, or R^2 and R^3 together with the nitrogen atom to which they are attached denote a heterocyclic group having 5 to 10 ring atoms of which 1, 2 or 3 are hetero atoms; R^4 and R^5 are each independently hydrogen or $\text{C}_1\text{-C}_6\text{-alkyl}$, or R^4 is hydrogen and R^5 is hydroxy- $\text{C}_1\text{-C}_6\text{-alkyl}$, acyl, $-\text{SO}_2\text{R}^6$ or $-\text{CON}(\text{R}^6)\text{R}^7$, or R^4 and R^5 together with the nitrogen atom to which they are attached denote a 5- or 6-membered heterocyclic group; R^6 and R^7 are each independently hydrogen or $\text{C}_1\text{-C}_6\text{-alkyl}$, or R^6 and R^7 together with the nitrogen atom to which they are attached denote a 5- or 6-membered heterocyclic group; and R^8 is $\text{C}_1\text{-C}_6\text{-alkyl}$, $\text{C}_1\text{-C}_6\text{-haloalkyl}$, or phenyl optionally substituted by $\text{C}_1\text{-C}_6\text{-alkyl}$.

Claim 12. (Currently Amended) A compound according to claim 11, which is

- (i) a compound of formula Ia or its pharmaceutically acceptable salts in free or salt form, wherein

Ar is phenyl substituted by halo;

X^1 is $-\text{S}-$, $-\text{S}(=\text{O})-$ or $-\text{S}(=\text{O})_2-$;

m is 2;

R^1 is $\text{C}_1\text{-C}_6\text{-alkyl}$ optionally substituted by hydroxy or $\text{C}_1\text{-C}_6\text{-alkoxy}$;

Y is oxygen;

R^2 is hydrogen; and

R^3 is a heterocyclic group having 5 to 11 ring atoms of which 1 to 4 are hetero atoms; or

- (ii) a compound of formula Ib or its pharmaceutically acceptable salts in free or salt form, wherein

Ar is phenyl substituted by halo;

X^2 is $-\text{O}-$, $-\text{C}(=\text{O})-$ or $-\text{CH}_2-$;

m is 1 or 2;

Q has the formula



where R^a is $\text{C}_1\text{-C}_6\text{-alkylene}$,

or Q is $-\text{C}(\text{R}^b)(\text{R}^c)-$ where R^b and R^c are independently $\text{C}_1\text{-C}_6\text{-alkyl}$

or R^b and R^c together form a $\text{C}_3\text{-C}_{10}\text{-cycloalkyl}$;

R^2 is hydrogen; and

R³ is a heterocyclic group having 5 to 11 ring atoms of which 1 to 4 are hetero atoms.

Claim 13. (Currently Amended) A compound according to claim 11, which is

(i) a compound of formula Ia or its pharmaceutically acceptable salts in free or salt form, wherein

Ar is phenyl substituted by halo, preferably chloro;

X¹ is -S-, -S(=O)- or -S(=O)₂-;

m is 2;

R¹ is C₁-C₄-alkyl optionally substituted by hydroxy or C₁-C₄-alkoxy;

Y is oxygen;

R² is hydrogen; and

R³ is a heterocyclic group having 5, 6 or 7 ring atoms of which one, two, three or four, are hetero atoms selected from nitrogen, oxygen and sulphur, said heterocyclic group being optionally substituted by C₁-C₄-alkyl, C₁-C₄-alkoxy or C₃-C₆-cycloalkyl; or

(ii) a compound of formula Ib or its pharmaceutically acceptable salts in free or salt form, wherein

Ar is phenyl substituted by halo, preferably chloro;

X² is -O-, -C(=O)- or -CH₂-;

m is 1 or 2;

Q has the formula



where R* is C₁-C₈-alkylene,

or Q is -C(R^b)(R^c)- where R^b and R^c are independently C₁-C₄-alkyl

or R^b and R^c together form a C₃-C₆-cycloalkyl;

R² is hydrogen; and

R³ is a heterocyclic group having 5, 6 or 7 ring atoms of which one, two, three or four, are hetero atoms selected from nitrogen, oxygen and sulphur, said heterocyclic group being optionally substituted by C₁-C₄-alkyl or C₃-C₆-cycloalkyl.

Claim 14. (Currently Amended) A compound according to claim 11 or a pharmaceutically acceptable salt thereof that is selected from the group consisting of:

1-((S)-3-[3-(4-Chloro-benzenesulfinyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(3,5-dimethoxy-phenyl)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfinyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-ethyl-[1,3,4]thiadiazol-2)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfinyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-ethyl-2-methyl-2H-pyrazol-3-yl)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfinyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-cyclopropyl-2-methyl-2H-pyrazol-3-yl)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfinyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-ethyl-isoxazol-3-yl)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfinyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(3-ethyl-isoxazol-5-yl)-urea;

1-((S)-3-[3-(4-Chloro-phenylsulfanyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-ethyl-[1,3,4]thiadiazol-2-yl)-urea;

1-((S)-3-[3-(4-Chloro-phenylsulfanyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-ethyl-2-methyl-2H-pyrazol-3-yl)-urea;

1-((S)-3-[3-(4-Chloro-phenylsulfanyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-cyclopropyl-2-methyl-2H-pyrazol-3-yl)-urea;

1-((S)-3-[3-(4-Chloro-phenylsulfanyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(3,5-dimethoxy-phenyl)-urea;

1-((S)-3-[3-(4-Chloro-phenylsulfanyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-ethyl-isoxazol-3-yl)-urea;

1-((S)-3-[3-(4-Chloro-phenylsulfanyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(3-ethyl-isoxazol-5-yl)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfonyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-ethyl-[1,3,4]thiadiazol-2-yl)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfonyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-ethyl-2-methyl-2H-pyrazol-3-yl)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfonyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-cyclopropyl-2-methyl-2H-pyrazol-3-yl)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfonyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(3,5-dimethoxy-phenyl)-urea;

1-((S)-3-[3-(4-Chloro-benzenesulfonyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(5-ethyl-isoxazol-3-yl)-urea; and

1-((S)-3-[3-(4-Chloro-benzene-sulfonyl)-azetidin-1-yl]-1-hydroxymethyl-propyl)-3-(3-ethyl-isoxazol-5-yl)-urea;

(+/-)-1-[(1R,2R)-2-[3-(4-Chloro-phenoxy)-azetidin-1-yl-methyl]-cyclohexyl]-3-(5-ethyl-
 [1,3,4]thiadiazol-2-yl)-urea;
 1-[(1R,2R)-2-[3-(4-Chloro-phenoxy)-azetidin-1-yl-methyl]-cyclohexyl]-3-(5-ethyl-2-methyl-2H-
 pyrazol-3-yl)-urea;
 1-[(1R,2R)-2-[3-(4-Chloro-phenoxy)-azetidin-1-yl-methyl]-cyclohexyl]-3-(5-cyclopropyl-2-methyl-
 2H-pyrazol-3-yl)-urea;
 1-[(1R,2R)-2-[3-(4-Chloro-phenoxy)-azetidin-1-yl-methyl]-cyclohexyl]-3-(5-cyclobutyl-2-methyl-
 2H-pyrazol-3-yl)-urea;
 1-[(1R,2R)-2-[3-(4-Chloro-phenoxy)-azetidin-1-yl-methyl]-cyclohexyl]-3-(2-ethyl-2H-tetrazol-5-
 yl)-urea;
 1-[(1R,2R)-2-[3-(4-Chloro-phenoxy)-azetidin-1-yl-methyl]-cyclohexyl]-3-(5-ethyl-isoxazol-3-yl)-
 urea;
 1-[(1R,2R)-2-[3-(4-Chloro-phenoxy)-azetidin-1-yl-methyl]-cyclohexyl]-3-(3-ethyl-isoxazol-5-yl)-
 urea;
 1-(1-[2-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-ethyl]-cyclobutyl)-3-(5-ethyl-[1,3,4]thiadiazol-2-yl)-
 urea;
 1-(1-[2-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-ethyl]-cyclobutyl)-3-(5-ethyl-2-methyl-2H-pyrazol-3-
 yl)-urea;
 1-(1-[2-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-ethyl]-cyclobutyl)-3-(5-cyclopropyl-2-methyl-2H-
 pyrazol-3-yl)-urea;
 1-(1-[2-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-ethyl]-cyclobutyl)-3-(5-cyclobutyl-2-methyl-2H-
 pyrazol-3-yl)-urea;
 1-(1-[2-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-ethyl]-cyclobutyl)-3-(2-ethyl-2H-tetrazol-5-yl)-urea;
 1-(1-[2-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-ethyl]-cyclobutyl)-3-(5-ethyl-isoxazol-3-yl)-urea;
 1-(1-[2-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-ethyl]-cyclobutyl)-3-(3-ethyl-isoxazol-5-yl)-urea;
 1-[3-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-1,1-dimethyl-propyl]-3-(5-ethyl-[1,3,4]thiadiazol-2-yl)-
 urea;
 1-[3-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-1,1-dimethyl-propyl]-3-(5-ethyl-2-methyl-2H-pyrazol-3-
 yl)-urea;
 1-[3-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-1,1-dimethyl-propyl]-3-(5-cyclopropyl-2-methyl-2H-
 pyrazol-3-yl)-urea;
 1-[3-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-1,1-dimethyl-propyl]-3-(5-cyclobutyl-2-methyl-2H-
 pyrazol-3-yl)-urea;
 1-[3-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-1,1-dimethyl-propyl]-3-(2-ethyl-2H-tetrazol-5-yl)-urea;
 1-[3-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-1,1-dimethyl-propyl]-3-(5-ethyl-isoxazol-3-yl)-urea; and
 1-[3-[3-(4-Chloro-phenoxy)-azetidin-1-yl]-1,1-dimethyl-propyl]-3-(3-ethyl-isoxazol-5-yl)-urea.

Claim 15. **(Currently Amended)** A pharmaceutical composition comprising a compound according to claim 11 or a pharmaceutically acceptable salt thereof in combination with another drug substance ~~which is selected from~~ an anti-inflammatory, a bronchodilator, an antihistamine or an anti-tussive substance.

Claim 16. **(Currently Amended)** A pharmaceutical composition comprising as active ingredient a compound according to claim 11, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier or diluent.

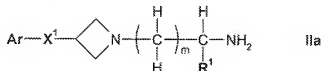
Claim 17. **(Currently Amended)** A pharmaceutical composition comprising as active ingredient a compound according to claim 14, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier or diluent.

Claim 18. **(Withdrawn – Currently Amended)**: A method of treating a condition mediated by CCR-3 in a subject in need of such treatment, which comprises administering to said subject an effective amount of a compound of formula I or a pharmaceutically acceptable salt thereof ~~as defined in claim 11 in free form or in the form of a pharmaceutically acceptable salt~~.

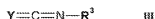
Claim 19. **(Withdrawn – Currently Amended)**: A method of treating an inflammatory or obstructive airways disease in a subject in need of such treatment, which comprises administering to said subject an effective amount of a compound of formula I or a pharmaceutically acceptable salt thereof ~~as defined in claim 11 in free form or in the form of a pharmaceutically acceptable salt~~.

Claim 20. **(Withdrawn)**: A process for the preparation of a compound of formula Ia or Ib as claimed in claim 11 which comprises

- (i) (A) for the preparation of compounds of formula Ia where R² is hydrogen, reacting a compound of formula IIa

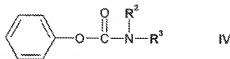


or a protected form thereof, where Ar, X¹, m and R¹ are as defined in claim 11, with a compound of formula III



where Y and R³ are as defined in claim 11; or

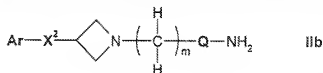
(B) for the preparation of compounds of formula Ia where Y is oxygen, reacting a compound of formula IIa where Ar, X¹, m and R¹ are as defined in claim 11, with a compound of formula IV



where R² and R³ are as defined in claim 11; or

(C) for the preparation of compounds of formula Ia where X¹ is -S(=O)₂-, oxidising a compound of formula Ia in protected form where X¹ is -S- and Ar, m, R¹, Y, R² and R³ are as defined in claim 11;

(D) for the preparation of compounds of formula Ib, reacting a compound of formula IIb



where Ar, X², m and Q are as defined in claim 11, with a compound of formula IV where R² and R³ are as defined in claim 11;

(E) for the preparation of compounds of formula Ib where R² is hydrogen, reacting a compound of formula IIb where Ar, X², m and Q are as defined in claim 11, with a compound of formula V



where R³ is as defined in claim 11; or

(F) for the preparation of compounds of formula Ib where X is -S(=O)₂-, oxidising a compound of formula Ib in protected form where X² is -S- and Ar, m, Q, R² and R³ are as defined in claim 11; and

(ii) recovering the product in free or salt form.